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MAY 1 9 2003

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<110> Azpiroz, Ricardo Choe, Sunghwa Feldmann, Kenneth A.

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TECH CENTER 1600/2900

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Pro Asp Ala Phe Ile Ala Ser Phe Val Arg Arg Phe Gly Arg Thr Gly
                                105
Val Tyr Arg Ser Phe Met Phe Ser Ser Pro Thr Val Leu Val Thr Thr
                            120
Ala Glu Gly Cys Lys Gln Val Leu Met Asp Asp Asp Ala Phe Val Thr
                        135
                                            140
Gly Trp Pro Lys Ala Thr Val Ala Leu Val Gly Pro Arg Ser Phe Val
                   150
                                        155
Ala Met Pro Tyr Asp Glu His Arg Arg Ile Arg Lys Leu Thr Ala Ala
                165
                                    170
Pro Ile Asn Gly Phe Asp Ala Leu Thr Gly Tyr Leu Pro Phe Ile Asp
                                185
Arg Thr Val Thr Ser Ser Leu Arg Ala Trp Ala Asp His Gly Gly Ser
                            200
Val Glu Phe Leu Thr Glu Leu Arg Arg Met Thr Phe Lys Ile Ile Val
                        215
Gln Ile Phe Leu Gly Gly Ala Asp Gln Ala Thr Thr Arg Ala Leu Glu
                    230
                                        235
Arg Ser Tyr Thr Glu Leu Asn Tyr Gly Met Arg Ala Met Ala Ile Asn
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Leu Pro Gly Phe Ala Tyr Arg Gly Ala Leu Arg Ala Arg Arg Arg Leu 265 260 Val Ala Val Leu Gln Gly Val Leu Asp Glu Arg Arg Ala Ala Arg Ala 280 Lys Gly Val Ser Gly Gly Gly Val Asp Met Met Asp Arg Leu Ile Glu 295 Ala Gln Asp Glu Arg Gly Arg His Leu Asp Asp Asp Glu Ile Ile Asp 310 315 Val Leu Val Met Tyr Leu Asn Ala Gly His Glu Ser Ser Gly His Ile 330 325 Thr Met Trp Ala Thr Val Phe Leu Gln Glu Asn Pro Asp Met Phe Ala 345 Arg Ala Lys Ala Glu Gln Glu Ala Ile Met Arg Ser Ile Pro Ser Ser 360 355 Gln Arg Gly Leu Thr Leu Arg Asp Phe Arg Lys Met Glu Tyr Leu Ser 375 380 Gln Val Ile Asp Glu Thr Leu Arg Leu Val Asn Ile Ser Phe Val Ser 395 390 Phe Arg Gln Ala Thr Arg Asp Val Phe Val Asn Gly Tyr Leu Ile Pro 405 410 Lys Gly Trp Lys Val Gln Leu Trp Tyr Arg Ser Val His Met Asp Pro 420 425 Gln Val Tyr Pro Asp Pro Thr Lys Phe Asp Pro Ser Arg Trp Glu Gly 440 435 His Ser Pro Arg Ala Gly Thr Phe Leu Ala Phe Gly Leu Gly Ala Arg Leu Cys Pro Gly Asn Asp Leu Ala Lys Leu Glu Ile Ser Val Phe Leu 475 470 His His Phe Leu Gly Tyr Lys Leu Ala Arg Thr Asn Pro Arg Cys 490 485 Arq Val Arq Tyr Leu Pro His Pro Arg Pro Val Asp Asn Cys Leu Ala 505 Lys Ile Thr Arg Val Gly Ser 515 <210> 23 <211> 492 <212> PRT <213> Danio rerio <400> 23 Met Gly Leu Tyr Thr Leu Met Val Thr Phe Leu Cys Thr Ile Val Leu Pro Val Leu Leu Phe Leu Ala Ala Val Lys Leu Trp Glu Met Leu Met 25 Ile Arg Arg Val Asp Pro Asn Cys Arg Ser Pro Leu Pro Pro Gly Thr 40 Met Gly Leu Pro Phe Ile Gly Glu Thr Leu Gln Leu Ile Leu Gln Arg 55 60 Arg Lys Phe Leu Arg Met Lys Arg Gln Lys Tyr Gly Cys Ile Tyr Lys 75 Thr His Leu Phe Gly Asn Pro Thr Val Arg Val Met Gly Ala Asp Asn 90 Val Arg Gln Ile Leu Leu Gly Glu His Lys Leu Val Ser Val Gln Trp 105 Pro Ala Ser Val Arg Thr Ile Leu Gly Ser Asp Thr Leu Ser Asn Val

His Gly Val Gln His Lys Asn Lys Lys Lys Ala Ile Met Arg Ala Phe 135 Ser Arg Asp Ala Leu Glu His Tyr Ile Pro Val Ile Gln Gln Glu Val 155 150 Lys Ser Ala Ile Gln Glu Trp Leu Gln Lys Asp Ser Cys Val Leu Val 170 165 Tyr Pro Glu Met Lys Lys Leu Met Phe Arg Ile Ala Met Arg Ile Leu 180 185 Leu Gly Phe Glu Pro Glu Gln Ile Lys Thr Asp Glu Gln Glu Leu Val 200 Glu Ala Phe Glu Glu Met Ile Lys Asn Leu Phe Ser Leu Pro Ile Asp 215 Val Pro Phe Ser Gly Leu Tyr Arg Gly Leu Arg Ala Arg Asn Phe Ile 230 235 His Ser Lys Ile Glu Glu Asn Ile Arg Lys Lys Ile Gln Asp Asp Asp 250 Asn Glu Asn Glu Gln Lys Tyr Lys Asp Ala Leu Gln Leu Leu Ile Glu 265 Asn Ser Arg Arg Ser Asp Glu Pro Phe Ser Leu Gln Ala Met Lys Glu 280 275 Ala Ala Thr Glu Leu Leu Phe Gly Gly His Glu Thr Thr Ala Ser Thr 295 300 Ala Thr Ser Leu Val Met Phe Leu Gly Leu Asn Thr Glu Val Val Gln 310 315 Lys Val Arg Glu Glu Val Gln Glu Lys Val Glu Met Gly Met Tyr Thr Pro Gly Lys Gly Leu Ser Met Glu Leu Leu Asp Gln Leu Lys Tyr Thr 345 Gly Cys Val Ile Lys Glu Thr Leu Arg Ile Asn Pro Pro Val Pro Gly 360 Gly Phe Arg Val Ala Leu Lys Thr Phe Glu Leu Asn Gly Tyr Gln Ile 375 Pro Lys Gly Trp Asn Val Ile Tyr Ser Ile Cys Asp Thr His Asp Val 390 395 Ala Asp Val Phe Pro Asn Lys Glu Glu Phe Gln Pro Glu Arg Phe Met 410 Ser Lys Gly Leu Glu Asp Gly Ser Arg Phe Asn Tyr Ile Pro Phe Gly 425 Gly Gly Ser Arg Met Cys Val Gly Lys Glu Phe Ala Lys Val Leu Leu 440 Lys Ile Phe Leu Val Glu Leu Thr Gln His Cys Asn Trp Ile Leu Ser 455 460 Asn Gly Pro Pro Thr Met Lys Thr Gly Pro Thr Ile Tyr Pro Val Asp 470 Asn Leu Pro Thr Lys Phe Thr Ser Tyr Val Arg Asn 485

<210> 24

<211> 504

<212> PRT

<213> Homo sapiens

<400> 24

Met Ala Leu Ile Pro Asp Leu Ala Met Glu Thr Trp Leu Leu Leu Ala 1 5 10 15

Val Ser Leu Val Leu Leu Tyr Leu Tyr Gly Thr His Ser His Gly Leu 20 25 30

Phe Lys Lys Leu Gly Ile Pro Gly Pro Thr Pro Leu Pro Phe Leu Gly Asn Ile Leu Ser Tyr His Lys Gly Phe Cys Met Phe Asp Met Glu Cys His Lys Lys Tyr Gly Lys Val Trp Gly Phe Tyr Asp Gly Gln Gln Pro Val Leu Ala Ile Thr Asp Pro Asp Met Ile Lys Leu Val Leu Val Lys Glu Cys Tyr Ser Val Phe Thr Asn Arg Glu Pro Phe Gly Pro Val Gly Phe Met Lys Ser Ala Ile Ser Ile Ala Glu Asp Glu Glu Trp Lys Arg Leu Arg Ser Leu Leu Ser Pro Thr Phe Thr Ser Gly Lys Leu Lys Glu Met Val Pro Ile Ile Ala Gln Tyr Gly Asp Val Leu Val Arg Asn Leu Arg Arg Glu Arg Glu Thr Gly Lys Pro Val Thr Leu Lys Asp Val Phe Gly Ala Tyr Ser Met Asp Val Ile Thr Ser Ser Ser Phe Gly Val Asn Val Asp Ser Leu Asn Asn Pro Gln Asp Pro Leu Val Glu Asn Thr Lys Lys Leu Leu Arg Phe Asp Phe Leu Asp Pro Phe Phe Leu Ser Ile Thr Val Phe Pro Phe Leu Ile Pro Ile Leu Glu Val Leu Asn Ile Cys Val Phe Pro Arg Glu Val Thr Asn Phe Leu Arg Lys Ala Val Lys Arg Met Lys Glu Ser Arg Leu Glu Asp Thr Gln Lys His Arg Val Asp Phe Leu Gln Leu Met Ile Asp Ser His Lys Asn Ser Lys Glu Thr Glu Ser His Lys Ala Leu Ser Asp Leu Glu Leu Val Ala Gln Ser Ile Ile Phe Ile Phe Ala Gly Tyr Glu Thr Thr Ser Ser Val Leu Ser Phe Ile Met Tyr Glu Leu Ala Thr His Pro Asp Val Gln Gln Lys Leu Gln Glu Glu Ile Asp Ala Val Leu Pro Asn Lys Ala Pro Pro Thr Tyr Asp Thr Val Leu Gln Met Glu Tyr Leu Asp Met Val Val Asn Glu Thr Leu Arg Leu Phe Pro Ile Ala Met Arg Leu Glu Arg Val Cys Lys Lys Asp Val Glu Ile Asn Gly Met Phe Ile Pro Lys Gly Trp Val Val Met Ile Pro Ser Tyr Ala Leu His Arg Asp Pro Lys Tyr Trp Thr Glu Pro Glu Lys Phe Leu Pro Glu Arg Phe Ser Lys Lys Asn Lys Asp Asn Ile Asp Pro Tyr Ile Tyr Thr Pro Phe Gly Ser Gly Pro Arg Asn Cys Ile Gly Met Arg Phe Ala Leu Met Asn Met Lys Leu Ala Leu Ile Arg Val Leu Gln Asn Phe Ser Phe Lys Pro Cys Lys Glu Thr Gln Ile Pro Leu Lys Leu Ser Leu Gly Gly Leu Leu Gln Pro Glu Lys Pro Val Val Leu Lys Val Glu Ser

495 485 490 Arg Asp Gly Thr Val Ser Gly Ala 500 <210> 25 <211> 575 <212> PRT <213> Artificial Sequence <220> <223> Consensus sequence <221> VARIANT <222> (1)...(575) <223> Xaa = Any Amino Acid or No Amino Acid <400> 25 25 Xaa Xaa Xaa Xaa Xaa Leu Leu Ser Xaa Xaa Ala Leu Xaa Val Xaa 40 Leu Xaa Leu Ala Ala Arg Arg Xaa Xaa Xaa Arg Tyr Xaa Xaa Xaa Xaa Xaa Xaa Xaa Arg Arg Lys Xaa Leu Pro Pro Gly Thr Met Gly Leu 70 75 Pro Xaa Leu Gly Glu Thr Leu Gln Phe Leu Lys Xaa Xaa Xaa Xaa Xaa 90 85 Xaa Pro Gly Asp Phe Xaa Lys Glu Arg Val Xaa Xaa Tyr Gly Xaa Xaa 100 105 Xaa Xaa Ile Tyr Lys His Leu Phe Gly Glu Pro Thr Ile Xaa Ser Xaa 120 125 115 Asp Ala Glu Leu Asn Arg Phe Xaa Leu Xaa Asn Glu Gly Xaa Lys Leu 135 Phe Xaa Cys Xaa Xaa Pro Ala Ser Xaa Xaa Gly Xaa Leu Gly Lys Xaa 150 Ser Leu Xaa Ala Xaa Xaa Gly Xaa Glu His Lys Arg Met Arg Xaa Leu 165 170 Leu Xaa Ser Xaa Phe Ser Xaa Xaa Xaa Leu Asp His Xaa Leu Pro 185 Xaa Ile Asp Arg Xaa Val Arg Ser Xaa Leu Xaa Xaa Trp Xaa Xaa Xaa 200 205 Xaa Gln Lys Xaa Xaa Ile Val Xaa Xaa Xaa Glu Xaa Lys Lys Met 215 220 Thr Phe Asp Xaa Xaa Xaa Lys Xaa Xaa Met Gly Xaa Xaa Pro Xaa Xaa 230 235 Glu Xaa Thr Xaa Xaa Xaa Leu Val Xaa Glu Xaa Glu Xaa Leu Ile 245 250 Lys Gly Leu Phe Ser Leu Pro Ile Asn Leu Pro Xaa Thr Ala Tyr Xaa Lys Ala Leu Xaa Ala Arg Ala Phe Xaa Xaa Ala Xaa Leu Glu Xaa Xaa 280 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Ile Xaa Glu Xaa Arg Xaa Glu Glu 295 320 310 315

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Xaa Xaa Xaa Xaa Xaa Asp Asp Leu Leu Gly Leu Leu Xaa Ala Xaa
               325
                                   330
Xaa Xaa Xaa Xaa Glu Asp Glu Xaa Xaa Xaa Leu Ser Asp Xaa
                               345
Glu Ile Xaa Asp Xaa Ile Xaa Xaa Leu Leu Phe Ala Gly His Glu Thr
                           360
Thr Ser Ser Xaa Leu Xaa Xaa Ala Val Lys Phe Leu Xaa Glu His Pro
                       375
                                           380
Asp Val Xaa Glu Xaa Leu Arg Glu Glu His Xaa Ala Ile Xaa Arg Ala
                   390
Lys Lys Xaa Xaa Xaa Glu Ser Xaa Leu Thr Xaa Xaa Asp Tyr Lys Lys
               405
                                    410
Met Xaa Tyr Thr Xaa Cys Val Ile Asn Glu Thr Leu Arg Leu Ala Xaa
                               425
           420
Ile Val Gly Gly Xaa Phe Arg Xaa Ala Xaa Lys Asp Val Glu Ile Asn
                           440
Gly Tyr Xaa Ile Pro Lys Gly Trp Lys Val Xaa Tyr Ser Ile Arg Ala
                       455
                                           460
Val His Leu Asp Pro Asp Xaa Tyr Pro Asp Pro Glu Lys Phe Asn Pro
                   470
                                       475
Xaa Arg Trp Xaa Xaa Lys Xaa Xaa Xaa Ser Asn Ser Xaa Xaa Xaa
               485
                                   490
Xaa Xaa Xaa Xaa Xaa Xaa Asn Xaa Asn Pro Phe Gly Gly Pro
                               505
Arg Leu Cys Pro Gly Lys Glu Leu Ala Lys Leu Glu Met Xaa Val Phe
                            520
Leu His Arg Leu Val Gln Xaa Phe Trp Glu Leu Ala Xaa Xaa Xaa Asp
                       535
                                           540
Xaa Xaa Xaa Lys Leu Val Xaa Phe Pro Thr Xaa Arg Pro Xaa Asp Asn
                   550
                                       555
Leu Pro Ile Lys Val Xaa Xaa Arg Asp Xaa Xaa Xaa Xaa Xaa
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<223> Heme binding domain
<221> VARIANT
<222> 4
<223> Xaa = Ala, Ser, or Val
<221> VARIANT
<222> 8
<223> Xaa = Any Amino Acid
<221> VARIANT
<222> 10
<223> Xaa = Pro, Ala, or Val
<400> 26
Pro Phe Gly Xaa Gly Arg Arg Xaa Cys Xaa Gly
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<223> Heme binding domain
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<211> 17
<212> PRT
<213> Artificial Sequence
<223> Signature sequence
<221> VARIANT
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Xaa Leu Leu Phe Ala Gly His Glu Thr Thr Ser Ser Xaa Ile Xaa Xaa
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1
Ala
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<213> Arabidopsis thaliana
<400> 30
Ala Gly His Glu Thr Ser
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